## WHAT IS CLAIMED IS:

1. A compound having the formula:
$P^{1}-X^{1}-(W^{1})_{n}-S-S-(W^{2})_{m}-X^{2}-P^{2}$ (I)
wherein
P1 and P2 are each members independently selected from the group consisting of a
hydrogen atom, an activating group and a protecting group;
$X^1$ and $X^2$ are each independently selected from the group consisting of a bond, -O-,
-NH-, -NR- and -CO <sub>2</sub> -, wherein R is a lower alkyl group having one to four
carbon atoms;
W <sup>1</sup> and W <sup>2</sup> are each independently selected from the group consisting of methylene,
oxyethylene and oxypropylene; and
n and m are each independently integers of from 2 to 12 with the proviso that n and
m are not the same when W1 and W2 are the same, and with the further
proviso that P <sup>1</sup> and P <sup>2</sup> are not both hydrogen atoms.
2. A compound in accordance with claim 1, wherein P <sup>2</sup> is an activating group
selected from the group consisting of a phosphoramidite, a trialkylammonium H-
phosphonate and a phosphate triester.
3. A compound in accordance with claim 1, wherein $P^2$ is a phosphoramidite,
P <sup>1</sup> is a protecting group selected from the group consisting of acid labile protecting groups,
$W^1$ and $W^2$ are both methylene, $X^1$ and $X^2$ are both -O-, and n and m are each integers of
from 2 to 8.
4. A compound in accordance with claim 1, wherein P <sup>2</sup> is a phosphoramidite,
P <sup>1</sup> is DMT, W <sup>1</sup> and W <sup>2</sup> are both methylene, X <sup>1</sup> and X <sup>2</sup> are both -O-, and n and m are each
integers of from 3 to 8.
5. A modified substrate for use in solid phase chemical synthesis, said substrate
having the formula:
$A^1-B^1-L^1 \tag{II}$

wherein  $A^1$  is a solid support,  $B^1$  is a bond or a derivatizing group, and  $L^1$  is a linking group 4 5 having the formula:  $P^1-X^1-(W^1)_n-S-S-(W^2)_m-X^2-$ 6 (IIa) 7 wherein, P<sup>1</sup> is a protecting group; 8  $X^1$  and  $X^2$  are each independently selected from the group consisting of a bond, -O-, 9 10 -NH-, -NR- and -CO<sub>2</sub>-, wherein R is a lower alkyl group having one to four 11 carbon atoms; W1 and W2 are each independently selected from the group consisting of methylene, 12 13 oxyethylene and oxypropylene; and 14 n and m are each independently integers of from 2 to 12 with the proviso that n and m are not the same when  $W^1$  and  $W^2$  are the same. 1 6. A substrate in accordance with claim 5, wherein P<sup>1</sup> is a photolabile protecting group. 1 7. A substrate in accordance with claim 5, wherein P<sup>1</sup> is a photolabile protecting group,  $W^1$  and  $W^2$  are both methylene, and  $X^1$  and  $X^2$  are both -O-. 1 8. A substrate in accordance with claim 5, wherein P<sup>1</sup> is a photolabile protecting group,  $X^1$  and  $X^2$  are both -O-, and n and m are each integers of from 2 to 8. 9. A substrate in accordance with claim 5, wherein P<sup>1</sup> is DMT, X<sup>1</sup> and X<sup>2</sup> are 1 both -O-, W1 and W2 are both methylene, and n and m are each integers of from 2 to 8. 1 10. A method of synthesizing small ligand molecules on a solid support having 2 optional spacers, said small ligand molecules being removable therefrom upon treatment 3 with a suitable disulfide cleaving reagent, said method comprising: 4 (a) contacting a solid support an unsymmetrical disulfide linking group of formula:  $P^1-X^1-(W^1)_n-S-S-(W^2)_m-X^2-P^2$ 5 (IIb) 6 wherein,

P<sup>1</sup> and P<sup>2</sup> are each members independently selected from the group consisting of a hydrogen atom, an activating group and a protecting group;

 $X^1$  and  $X^2$  are each independently selected from the group consisting of a bond, -O-, -NH-, -NR- and -CO<sub>2</sub>-, wherein R is a lower alkyl group having one to four carbon atoms;

 $W^1$  and  $W^2$  are each independently selected from the group consisting of methylene, oxyethylene and oxypropylene; and

n and m are each independently integers of from 2 to 12 with the proviso that n and m are not the same when  $W^1$  and  $W^2$  are the same;

to produce a derivatized solid support having attached unsymmetrical disulfide linking groups suitably protected with protecting groups;

- (b) optionally removing said protecting groups from said derivatized solid support to provide a derivatized solid support having unsymmetrical disulfide linking groups with synthesis initiation sites; and
- (c) coupling said small ligand molecules to said synthesis initiation sites on said derivatized solid support to produce a solid support having attached small ligand molecules which are removable therefrom upon application of said disulfide cleaving reagent.

## 11. A compound of the formula:

$$\begin{array}{c} P^{11}O \\ OP^{12} \\ H_3C \\ CH_3 \end{array} \qquad \begin{array}{c} H \\ O \\ CH_3 \end{array} \qquad \begin{array}{c} O \\ CH_3 \\ CH_3 \end{array} \qquad \begin{array}{c} (VI) \\ CH_3 \\ \end{array}$$

wherein P<sup>11</sup> and P<sup>12</sup> are each independently selected from the group consisting of hydrogen, a protecting group, and a phosphodiester-forming group.

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- 1 12. A compound in accordance with claim 11, wherein  $P^{11}$  and  $P^{12}$  are both hydrogen.
- 1 13. A compound in accordance with claim 11, wherein  $P^{11}$  is a protecting group and  $P^{12}$  is a phosphoramidite.
- 1 **14.** A compound in accordance with claim 11, wherein  $P^{11}$  is DMT and  $P^{12}$  is a phosphoramidite.
- 1 15. A substrate for the solid phase synthesis of oligonucleotides, said substrate 2 having the formula:
- $A^{11}$ - $B^{11}$ - $L^{11}$ -FI
- 4 wherein A<sup>11</sup> is a solid support, B<sup>11</sup> is a bond or a derivatizing group, L<sup>11</sup> is a linking group,
- 5 and Fl is a fluorescent moiety having the formula:

$$\begin{array}{c} P^{11}O \\ OP^{12} \\ H_3C \\ CH_3 \end{array}$$
 (VI)

- 7 wherein one of  $P^{11}$  and  $P^{12}$  is a covalent bond to  $L^{11}$  and the other of  $P^{11}$  and  $P^{12}$  is selected from the group consisting of hydrogen, a protecting group, and a phosphoramidite.
- 1 16. A substrate bound, fluorescently labeled oligonucleotide having the formula:
- 2 A<sup>11</sup>-B<sup>11</sup>-L<sup>11</sup>-Nu-Fl
- 3 wherein A<sup>11</sup> is a solid support, B<sup>11</sup> is a bond or a derivatizing group, L<sup>11</sup> is a linking group,
- 4 Nu is an oligonucleotide and Fl is a fluorescent moiety having the formula:

wherein one of  $P^{11}$  and  $P^{12}$  is a covalent bond to  $L^{11}$  and the other of  $P^{11}$  and  $P^{12}$  is selected from the group consisting of hydrogen, a protecting group, and a phosphoramidite.

1 17. A substrate bound, fluorescently labeled oligonucleotide having the formula:

2 
$$A^{11}$$
- $B^{11}$ - $L^{11}$ - $F$ l- $Nu$ 

- 3 wherein A<sup>11</sup> is a solid support, B<sup>11</sup> is a bond or a derivatizing group, L<sup>11</sup> is a linking group,
- 4 Fl is a fluorescent moiety having the formula:

wherein each of  $P^{11}$  and  $P^{12}$  represents a bond; and Nu is an oligonucleotide.

- 1 18. A selectively cleavable linkage molecule useful in solid phase compound
- 2 synthesis, said linkage molecule having the formula:

1

2

1

2

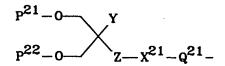
3

 $p^{21}-0$  Y  $P^{22}-0$   $Z-X^{21}-0$ 

4 wherein P<sup>21</sup> and P<sup>22</sup> are each protecting groups with the provisos that P<sup>21</sup> can be removed 5 under conditions which will not remove P<sup>22</sup>, and P<sup>22</sup> can be removed under 6 conditions which will not remove P21; 7 X<sup>21</sup> is a linking moiety selected from the group consisting of an alkylene chain and an 8 9 aryl group; Y is a substituent selected from the group consisting of -C(=O)R, -S(O)R, -S(O)R, 10 -S(O)<sub>2</sub>NRR', -CN, -CF<sub>3</sub>, -NO<sub>2</sub> and a phenyl ring having one or more 11 12 substituents selected from the group consisting of halogen, nitro, cyano and 13 trifluoromethyl; Z is a linking moiety selected from the group consisting of -C(=O), -S(O), 14  $-S(O)_{2}-, -S(O)_{2}NR-,$ 15 16 wherein 17 R and R' are each independently selected from the group consisting of hydrogen, C<sub>1</sub>-C<sub>12</sub> alkyl and aryl; and 18 Q is a phosphate ester-forming group selected from the group consisting of a 19 phosphoramidite and a trialkylammonium H-phosphonate.

- 19. A selectively cleavable linkage molecule in accordance with claim 18, wherein  $X^{21}$  is an amino alkoxy group, Y is -C(=O)R, Z is -C(O)— and Q is a phosphoramidite.
- 20. A selectively cleavable linkage molecule in accordance with claim 18, wherein  $P^{21}$  is removable under photolytic conditions,  $P^{22}$  is removable under acidic conditions,  $X^{21}$  is an amino alkoxy group, Y is -C(=O)R, Z is -C(O) and Q is a phosphoramidite.

- 21. A selectively cleavable linkage molecule in accordance with claim 18, wherein P<sup>21</sup> is MeNPOC, P<sup>22</sup> is DMT, X<sup>21</sup> is -NH-CH<sub>2</sub>CH(CH<sub>2</sub>)-O-, Y is -C(=O)R, Z is
- wherein  $P^{21}$  is MeNPOC,  $P^{22}$  is DMT,  $X^{21}$  is -NH-CH<sub>2</sub>CH(CH<sub>3</sub>)-O-, Y is -C(=0)R -C(O)- and O is a phosphoramidite.
- 22. A modified substrate for use in solid phase chemical synthesis, said substrate having the formula:
- $L^{21}$ - $B^{21}$ - $A^{21}$
- 4 wherein  $A^{21}$  is a solid support,  $B^{21}$  is a bond or a derivatizing group, and  $L^{21}$  is a linking
- 5 group having the formula:



7 wherein

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- 8  $P^{21}$  and  $P^{22}$  are each protecting groups with the provisos that  $P^{21}$  can be removed
- 9 under conditions which will not remove P<sup>22</sup>, and P<sup>22</sup> can be removed under
- 10 conditions which will not remove P<sup>21</sup>;
- $X^{21}$  is a linking moiety selected from the group consisting of an alkylene chain and an
- 12 aryl group;
- Y is a substituent selected from the group consisting of -C(=O)R, -S(O)R, -S(O)R,
- -S(O)<sub>2</sub>NRR', -CN, -CF<sub>3</sub>, -NO<sub>2</sub> and a phenyl ring having one or more
- substituents selected from the group consisting of halogen, nitro, cyano and
- trifluoromethyl;
- Z is a linking moiety selected from the group consisting of -C(=O), -S(O),
- 18  $-S(O)_2$ -,  $-S(O)_2NR$ -,
- wherein
- 20 R and R' are each independently selected from the group consisting of
- 21 hydrogen, C<sub>1</sub>-C<sub>12</sub> alkyl and aryl; and
  - Q<sup>21</sup> is a phosphate ester linking group.